

In the specification

Please replace the paragraph on page 35, lines 8-31 with the following paragraph.

The cloned p35 cDNA, after sequence confirmation, was adapted for expression as a fusion protein as follows. At the fusion junction, the C-terminal amino acid residue of the CH3 is lysine and the N-terminal residue of the mature p35 is arginine. To minimise proteolysis at the fusion junction with these two basic residues, both of them were mutagenised to alanine, which, in the case of IL2 immunocytokine, has been shown to extend serum half-life [Gillies *et al.* (2002) *Clin. Cancer Res.* 8:210]. For reconstruction of the fusion junction, there is a convenient *Bal*I site just 11 base-pairs (bp) downstream of the mature N-terminus of p35. Hence a *Xma*I-*Bal*I oligonucleotide linker consisting of sense strand 5'-CCG GGC **GCC GCA** AAC CTC CCC GTG G-3' [SEQ ID NO:22] and anti-sense strand 5'- C CAC GGG GAG GTT **TGC GGC GC**-3' [SEQ ID NO:23], where the **GCC GCA** [SEQ ID NO: 24] denote the two alanine substitutions, was synthesised and ligated to a *Bal*I-*Xho*I restriction fragment encoding the rest of the p35 subunit. The resultant *Xma*I-*Xho*I fragment in turn was ligated to the unique *Xma*I site in the pdHL11 expression vector, forming the CH3-p35 fusion junction. The peptide sequence at the junction, LSLSPGAANLPV [SEQ ID NO: 25] [~~SEQ ID NO:24~~], where **AA** are the two alanine substitutions, is novel and potentially immunogenic. Indeed it contained a potential T helper cell epitope, which could be removed by mutating the LSLS [SEQ ID NO: 26] residues to ATAT [SEQ ID NO: 27], based on Biovation's technology called deImmunization. The resultant deImmunised fusion junction sequence is called M1. Therefore, the huBC1-H chain-M1-hu p35 DNA consists of the *Xho*I-HindIII fragment encoding the signal peptide-VH,

the HindIII-XmaI fragment encoding the genomic human IgG1 H chain constant regions with the deImmunised junction, and the XmaI-XhoI fragment encoding the p35 subunit.

Please replace the paragraph on page 46, 1-2 with the following paragraph.

PEHFSGRPREDRVPHSRNSITLTNLTPGTEYVVSIVAL

NGREESPLLIGRSRSHHHHHH"

[SEQ ID NO: 28]~~[SEQ ID~~

~~NO: 25]~~

Please replace the paragraph on page 46, lines 5-9 with the following paragraph.

Note1: Residue 1 to 207 is pQE sequence from and including Qiagen promoter primer (CCCGAAAAGTGCCACCTG) **[SEQ ID NO:29]**. Residue 1069 to 1126 is pQE12 sequence from the end of the hexa-histidine tag to the Qiagen reverse primer sequence (GTTCTGAGGTCATTACTGG) **[SEQ ID NO:30]**. Fibronectin-derived sequence (*i.e.* without MRGS and hexa-histidine tag is in lower case).

Please replace the paragraph on page 46, line 18 until page 47, line 4 with the following paragraph.

1 CCCCGAAAAG TGCCACCTGA CGTCTAAGAA ACCATTATTA TCATGACATT
AACCTATAAA

61 AATAGGCGTA TCACGAGGCC CTTTCGTCTT CACCTCGAGA AATCATAAAA
AATTTATTTG

121 CTTTGTGAGC GGATAACAAT TATAATAGAT TCAATTGTGA GCGGATAACA
ATTCACACA

181 GAATTCATTA AAGAGGAGAA ATTA ACTATG AGAGGATCtg tggtagacacc
attgtctcca

241 ccaacaaact tgcactctgga ggcaaacct gacactggag tgctcacagt ctctgggag

301 aggagcacca cccagacat tactgggtat agaattacca caaccctac aaacggccag

361 cagggaaatt ctttgaaga agtgggtccat gctgacaga gctctgcac tttgataac

421 ctgagtcctg gcctggagta caatgtcagt gtttactctg tcaaggatga caaggaaagt

481 gtccttatct ctgataccat catccagct gttctctc cactgacct gcgattcacc

541 aacattggc cagacacat gcgtgtcacc tgggtccac cccatccat tgatttaacc

601 aacttctgg tgcgttactc acctgtgaaa atgaggaag atgttcaga gttgtcaatt

661 tctcttcag acaatgcagt ggtcttaaca aatctctgc ctggtacaga atatgtagt

721 agtgtctcca gtgtctacga acaacatgag agcacacctc ttagaggaag acagaaaaca

781 ggtcttgatt cccaactgg cattgacttt tctgatatta ctgccaactc tttactgtg

841 cactggattg ctctcgagc caccatcact ggctacagga tccgcatca tcccgagcac

901 ttcagtggga gacctcgaga agatcgggtg cccactctc ggaattccat caccctcacc

961 aacctcactc caggcacaga gtatgtggtc agcatcgttg ctctaatgg cagagaggaa

1021 agtccttat tgattggcaG ATCCAGATCT CATCACCATC ACCATCACTA

AGCTTAATTA

1081 GCTGAGCTTG GACTCCTGTT GATAGATCCA GTAATGACCT CAGAAC

[SEQ ID NO: 31] ~~[SEQ ID NO: 26]~~

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Please replace the paragraph on page 47, lines 6-36, with the following paragraph.

(b) *Fibronectin 7B89 fragment*

LOCUS FN7B89.DNA 1399 bp mRNA PRI 01-OCT-1999

DEFINITION Human mRNA for fibronectin domains 7B89 in pQE12 (pAS33)

NID Derived from g31396 and pQE12 (Qiagen).

VERSION X02761.1 GI:31396

KEYWORDS alternate splicing; fibronectin.

SOURCE human.

ORGANISM Homo sapiens

Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Mammalia;

Eutheria; Primates; Catarrhini; Hominidae; Homo.

CDS <208..1341

/product="Fn MRGS-7B89-HHHHHH"

/translation="

MRGSVVTPLSPPTNLHLEANPDTGVLTVSWERSTTPDI

TGYRITTTPTNGQQGNSLEEVVHADQSSCTFDNLSPGL

EYNVSVYTVKDDKESVPISDTIPEVPQLTDLSFVDIT

DSSIGLRWTPLNSSTIIGYRITVVAAGEGIPIFEDFVD

SSVGYYTVTGLEPGIDYDISVITLINGGESAPTTLTQQ

TAVPPPTDLRFTNIGPDTMRVTWAPPPSIDLTNFLVRY

SPVKNEEDVAELSISPSDNAVVLTNLLPGTEYVVS

VYEQHESTPLRGRQKTGLDSPTGIDFSDITANSFTVHW

IAPRATITGYRIRHHPEHFSGRPREDRVPHSRNSITLT

NLTPGTEYVVSIVALNGREESPLLIGRSRSHHHHHH"

[SEQ ID NO:32] ~~**[SEQ ID NO: 27]**~~

Note1: Residue 1 to 207 is pQE sequence from and including Qiagen promoter primer (CCCGAAAAGTGCCACCTG) [SEQ ID N:29]. Residue 1342 to 1399 is pQE12 sequence from the end of the hexa-histidine tag to the Qiagen reverse primer sequence (GTTCTGAGGTCATTACTGG) [SEQ ID NO:30]. Fibronectin-derived sequence (*i.e.* without MRGS and hexa-histidine tag is in lower case).

Please replace the paragraph on page 48, lines 5-38 with the following paragraph.

BASE COUNT 390 a 368 c 290 g 351 t
ORIGIN

1 CCCCGAAAAG TGCCACCTGA CGTCTAAGAA ACCATTATTA TCATGACATT
AACCTATAAA

61 AATAGGCGTA TCACGAGGCC CTTTCGTCTT CACCTCGAGA AATCATAAAA
AATTTATTTG

121 CTTTGTGAGC GGATAACAAT TATAATAGAT TCAATTGTGA GCGGATAACA
ATTCACACA

181 GAATTCATTA AAGAGGAGAA ATTA ACTATG AGAGGATCtg tggtgacacc
attgtctcca

241 ccaacaaact tgcacttgga ggcaaacct gacactggag tgctcacagt ctctggggag

301 aggagcacca cccagacat tactgggttat agaattacca caaccctac aaacggccag

361 cagggaaatt ctttgaaga agtgggccat gctgatcaga gctcctgcac tttgataac
421 ctgagtcccg gcctggagta caatgtcagt gtttacctg tcaaggatga caaggaaagt
481 gtccttatct ctgataccat catcccagag gtgcccacac tcaactgacct aagctttgtt
541 gatataaccg attcaagcat cggcctgagg tggaccccg taaactcttc caccattatt
601 gggtagcgca tcacagtagt tgcggcagga gaaggtatcc ctattttga agattttgtg
661 gactcctcag taggatacta cacagtcaca gggctggagc cgggcattga ctatgatata
721 agcggtatca ctctcattaa tggcggcgag agtgccccta ctacactgac acaacaaacg
781 gctgttctc ctcccactga cctgcgattc accaacattg gtccagacac catgcgtgtc
841 acctgggctc cccccctc cattgattta accaacttc tggcggtta ctacactgtg
901 aaaaatgagg aagatgttgc agagttgtca atttctctt cagacaatgc agtggctta
961 acaaatctcc tgcctgttac agaatatga gtgagtgtct ccagtgtcta cgaacaacat
1021 gagagcacac ctcttagagg aagacagaaa acaggtcttg attccccaac tggcattgac
1081 tttctgata ttactgcaa ctctttact gtgcactgga ttgctctcg agccaccatc
1141 actggctaca ggatccgcca tcatcccgag cacttcagtg ggagacctcg agaagatcgg
1201 gtgcccact ctggaattc catcacctc accaacctca ctccaggcac agagtatgtg
1261 gtcagcatcg ttgctcttaa tggcagagag gaaagtcctt tattgattgg caGATCCAGA
1321 TCTCATCACC ATCACCATCA CTAAGCTTAA TTAGCTGAGC TTGGACTCCT
GTTGATAGAT
1381 CCAGTAATGA CCTCAGAAC

[SEQ ID NO:33] [SEQ ID NO: 28]

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